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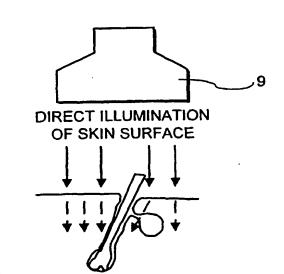
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(54) Title: COSMETIC DEPILATION KIT WITH PHOTOSENSITIZER



(57) Abstract: A marker substance includes a photosensitive toxic material to tag subcutaneous hair growth cells of a target zone of skin. Illumination apparatus is used to illuminate the skin in the target zone with light of a wavelength selected to effect preferential interaction with the hair growth cells tagged with the marker substance, resulting in a toxic effect inhibiting the hair growth capability of the tagged illuminated cells; wherein. In the kit the marker substance may be provided in a form comprising discrete portions of the photosensitive toxic material and a carrier material, the portions to be mixed prior to application to the target zone. Additionally or alternatively, the illumination apparatus includes an applicator configured to deliver non-laser light and/or configured to be placed against the target zone skin to deliver the illumination to the tagged hair growth cells.

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COSMETIC DEPILATION KIT WITH PHOTOSENSITIZER

The present invention relates to a cosmetic depilation kit.

Laser hair removal is currently an effective and long term method for effecting cosmetic hair removal. An exemplary laser depilation technique is disclosed in EP-A-0732895. In such techniques, high intensity laser energy is directed towards the skin selectively heating hair surrounding the follicle leading to damage of the hair growth mechanism. If sufficient damage is induced, the hair follicle is effectively destroyed and the hair will not regrow. This technique has disadvantages, one such being that scarring and pigmentary changes can occur. This is particularly the case where the individual being treated has dark skin and/or light hair. Such techniques are also relatively slow, with the target zone needing to be covered by repeated laser pulses.

US-A-5669916 discloses a technique in which a hair is first plucked from a follicle before a photosensitive toxic material is directed along the path of the removed hair to the follicle. The toxic material is then activated by irradiation using laser.

An improved technique has now been devised.

According to a first aspect, the present invention provides a cosmetic kit for use in inhibiting hair growth, the kit comprising:

- a) a marker substance including a photosensitive toxic material to tag subcutaneous hair growth cells of a target zone of skin; and,
- illumination apparatus for illuminating the skin in the target zone with light of a wavelength selected to effect preferential interaction with the hair growth cells tagged with the marker substance, resulting in a toxic effect inhibiting the hair growth capability of the tagged illuminated cells; wherein:
 - (i) the marker substance is provided in a form comprising discrete portions of the photosensitive toxic material and a carrier material, the portions to be mixed prior to application to the target zone; and or,
 - (ii) the illumination apparatus includes an applicator configured to deliver non-laser light and/or configured to be placed against the target zone skin to deliver the illumination to the tagged hair growth cells.
- The photosensitive toxic material is a material whose toxicity increases substantially upon exposure

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illumination of predetermined wavelength at sufficient energy delivered. One such material is ALA (aminolevulinic acid); this and other materials are known in the art and described, for example in US-A-5669916.

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The carrier material is beneficially a material easily an preferably absorbed into the skin comprises emollient moisturiser or absorbent carrying the photosensitive toxic material in dissolved form or It is important to maintain the carrier and otherwise. the photosensitive toxic material separate until shortly prior to use in order to inhibit any adverse interaction between the materials prior to application to the skin of the target zone.

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The portions of the photosensitive toxic material and the carrier material are dosed in a predetermined ratio. In one embodiment the proportion would be approximately 25% photosensitive toxic material to approximately 75% carrier material.

20 material

The kit beneficially includes a dispenser or a containerdispenser for dispensing the marker substance.

A container for the carrier material and or the photosensitive toxic material preferably affords light shielding of the photosensitive toxic material.

Mixing means is beneficially provided for mixing the photosensitive toxic material and the carrier material.

The mixing means preferably permits mixing of the

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photosensitive toxic material and the carrier material at a zone substantially sealed from the user.

Beneficially a container for the photosensitive toxic material and the carrier material comprises respective sealed zones containing the photosensitive toxic material and the carrier material respectively, the zones being configured to be selectively breached or ruptured permitting mixing of the materials. A mixing zone may be provided adjacent the breachable sealed zones, the mixing zone preferably communicating with a dispensing outlet.

The illumination apparatus preferably includes window means varranged to overlay the target zone skin, the illumination being directed via the window means.

In one embodiment, the illumination apparatus includes an illumination applicator having a flexible structure arranged to conform to the target zone skin body part.

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The illumination apparatus preferably includes:

 an illumination applicator structure arranged to deliver illumination over a predetermined area;

- b) a waveguide to connect with the illumination applicator; and,
- c) a light source to direct light along the waveguide.

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The illumination apparatus beneficially includes an illumination applicator structure having internal light reflecting walls and an associated light transmissive zone means through which reflected light is permitted to pass. This may be achieved, for example, where the applicator structure comprises one or more lengths of fibreoptic (preferably the applicator structure comprises a plurality of lengths of fibre optic, the lengths being arranged to deliver light at different zones of the applicator structure).

The marker substance is desirably preferentially absorbed by, or tagged to, hair growth cells in the target zone. The excess proportion of the applied marker substance not preferably the hair growth cells is absorbed bv metabolised and removed from the tissue by the normal action of the body. The technique according to the invention has been found to be effective even without the prior mechanical/physical depilation suggested in US-A-Indeed such a step is discouraged due to the unnecessary trauma and pain it entails.

The hair growth cells targeted for tagging are preferably so called 'stem' cells in the 'bulge' area of the follicle (adjacent the sebaceous gland). By effectively destroying or disabling the 'stem' cells once the hair has completed a growth cycle, a new hair cycle will be inhibited from starting, thereby preventing the follicle from functioning properly. The marker substance is therefore selected to preferentially tag 'stem' cells.

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The marker substance is preferably topically applied to the skin at the target zone. The marker substance penetrates into the skin to preferentially tag the target cells (typically by being preferentially absorbed). The marker substance preferably comprises a photo therapy drug such as for example 5-Aminolaevolinic Acid (5-ALA). Desirably the drug is provided in a topically administrable form such as a cream or the like.

The interaction of the illuminating light with the marking substance preferably results in the desired hair growth inhibition. The interaction may be photochemical and/or photo-thermal in nature. For photochemical interaction, light of a predetermined wavelength induces a chemical reaction between the marker substance and the tagged cells resulting in destruction of, or disabling of, the tagged cells. For photo-thermal interaction, the incident light causes a temperature rise in the tagged cells to a degree sufficient to destroy or disable the tagged cell.

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The wavelength of the illuminating light is preferably selected such that it is absorbed by the marker substance (or the tagged cell). The wavelength of the illuminating light is preferably substantially matched to the optical absorption characteristics of the marker substance.

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The illuminating light is preferably substantially in the range 500nm-1000nm wavelength. The light is preferably concentrated within a relatively narrow bandwidth within the range. This may be achieved by filtering of the light or use of light emitting apparatus arranged to emit a

discrete wavelength (or band of wavelengths). For example where 5-ALA is used as the marker substance to tag the cells it is preferred that the illuminating light includes on or more wavelengths substantially in the range 600nm-650nm.

The illuminating light may, for example, be laser (including semiconductor laser), white light (preferably filtered), or LED light.

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The intensity of the light is determined by the quantity of light required to produce the necessary interaction with the tagged cells. Generally however the intensity required will be substantially lower than the intensity required for prior art laser depilation techniques. This enables the technique to be more suitable for unsupervised 'Home' use.

The light may be pulsed or applied as continuous wave. The light typically floods the target zone, which target zone is typically of an area in the range 0.5cm²-10cm², more preferably 1cm²-5cm². The energy applied per pulse to the target zone is preferably substantially in the range 40J or less (more preferably 25J or less).

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The possibility of using lower intensity and non-laser light also enables a greater area target zone to be simultaneously illuminated, with reduced risk of skin burn. The technique is therefore safer and quicker.

The invention will now be further described, in a specific embodiment, by way of example only and with reference to the accompanying drawings in which:

Figure 1 is a schematic representation of a mammalian hair follicle:

Figure 2 is a schematic representation, similar to Figure 1, showing a first stage in application of the depilation technique;

Figure 3 is a schematic representation similar to Figures 1 and 2, showing a subsequent stage in the technique;

15 Figures 4 and 5 are schematic plan and side views respectively of a container-dispenser for the marker substance according to the invention;

Figures 6 and 7 show alternative general embodiments of illumination apparatus for inclusion in a kit according to the invention;

Figure 8 is a schematic view of a laser diode illumination apparatus for inclusion in a kit according to the invention;

Figure 9 is a schematic view of LED illumination apparatus for inclusion in a kit according to the invention;

Figure 10 is a schematic view of 'white light' illumination apparatus for inclusion in a kit according to

the invention;

Figure 11 is a schematic view of alternative 'white light' illumination apparatus for inclusion in a kit according to the invention;

Figure 12 is a schematic view of a pad or patch applicator structure illumination apparatus for inclusion in a kit according to the invention;

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Figure 13 is a perspective view of the applicator pad or patch of figure 12;

Figure 14 is a sectional view of the fibreoptic used in the pad or patch of figures 12 and 13; and

Figure 15 is a schematic longitudinal section of the fibreoptic of figure 14.

Referring to the drawings and initially to Figure 1, there is shown human hair follicle supporting a hair shaft 1 growing through the epidermis 2 and dermis 3 of mammalian skin tissue. 'Stem' cells 7 are present in the 'bulge' region 4 adjacent the sebaceous gland 5. Stem cells have a relatively high turnover/life cycle and are responsible for instituting cyclical growth of the hair from the follicle.

In accordance with the invention, a drug formulation including 5-Aminolaevolinic Acid (5-ALA) is provided in a form for topical administration to the outer surface of

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the skin 6. The drug 5-ALA is selected because it is selectively absorbed and retained in stem cells 7 when penetrating into the epidermal layer 3 (via epidermis 2) thereby effectively tagging the stem cells. Concentrations of the drug not absorbed in the stem cells 7 are metabolised and removed from the tissue by the normal action of the body.

After a predetermining period of time to allow the excess "marker" drug to be removed from the target zone tissue (other than from the "tag" stem cells 7), the target zone of the skin is directly illuminated from externally of the body by means of an illumination device 9 which supplies light of a preselected wavelength matched to a preselected characteristic absorption wavelength of the marker drug (now tagging stem cells 7).

The intensity of the light supplied, and duration of supply of the light, is controlled such that either photochemical or photo-thermal interaction (or both) of the light with the tagged stem cells 7 causes disabling or destruction of the relevant stem cell 7 without disabling or destroying, substantially, the surrounding tissue material. This results in the tagged and disabled stem cell being inhibited from acting further to produce hair from the relevant follicle.

The technique, by selectively marking/tagging and destroying stem cells preferentially (and the associated matching of the wavelength of the illuminating light) enables relatively low intensity light to be used, which

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may flood the target zone of the skin enabling relatively large area of hair covered skin to be treated liuhi simultaneously. Typically, ille illumination device is pulsed to 9 enable thermal relaxation of the skin to occur between consecutive Typically, the energy supplied to the skin pulses. surface by the illuminating device 9 does not exceed 25J, before significant thermal relaxation of the skin is permitted.

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The illuminating device may comprise laser, semiconductor laser, white light illumination device (typically with a filter) or an LED device. The wavelength band width emitted where 5-ALA is used as the marker is preferably substantially within the range 600nm - 650nm (which may be achieved by filtering if required).

The low intensity nature of the light required to effect depilation using the technique according to the invention, enables the technique to be relatively safe when compared to prior art laser hair removal techniques. The drug formulation and illumination device may therefore be sold as a consumer kit for "home" use.

Referring to figures 4 to 15 a "home use" kit for cosmetic depilation would typically comprise a power supply and lighting apparatus 10, 11 (see figures 6 and 7) for direct connection to mains power supply. The apparatus 10, 11 either includes flexible electric connection to a light source 12 for application of illuminating radiation to the skin or, a "fixed" light source 13 and fibreoptic

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connection 14 to an applicator 16.

The other essential element of the kit is the market substance typically provided in a container 20 (see figures 4 and 5) which comprises a heat sealed plastics container/dispenser 20 having sealed chambers 21, 22, 23.

Chamber 21 contains the ALA material and chamber 23 carries the moisturiser/emollient carrier cream. A manual pinch pressure causes seals 24, 25 to be ruptured and the respective materials to be urged from chambers 21, 23 into mixing chamber 22 where they become thoroughly, and intimately, mixed. A cap 26 is subsequently broken from the end of nozzle 27 permitting the ALA and carrier mixture to be squeezed out of the container dispenser 20 under manual pressure.

The container dispenser 20 ensures that the correct proportions of ALA drug and carrier are mixed, and that mixing may be delayed until the point of application.

Following application of the mixture topically to the target zone of skin, and subsequently waiting the required length of time to permit the concentration of ALA in the non-target tissue to be reduced to the required low level, the illumination apparatus is applied to the skin and switched on.

Various forms of illuminating apparatus may be used. The
apparatus 110 in figure 8 includes a laser diode light
source 125 contained within a protective casing 126 and

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including focussing optics 127 arranged to ensure the light is directed through a protective window 128. Typically the window 128 is held against the skin of the target zone to ensure correct delivery of the illuminating light to the target zone.

In the embodiment shown in figure 9 a 2D array of LEDs (light emitting diodes) 225 is utilised. Focussing optics 227 ensure the light is directed through the protective window 228 in a similar manner to the arrangement of figure 8.

In the apparatus shown in figure 10, an electric filament "white light" source is used. The source 325 is surrounded by a protective housing 326 including a shaped reflective surface 327 arranged to direct the light out of the housing via protective window 328.

In the embodiment shown in figure 11, a white light filament source 325 is once again used. In this embodiment an internally reflecting light guide (such as a fibreoptic) is used. The fibreoptic may be connected to a light applicator to be connected to the skin as will be described hereafter.

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The apparatus of figure 12 shows an applicator pad or patch structure 426 which is typically flexible in nature, permitting the pad 426 to be conformed to the shape of the body part to which it is applied. As shown in figures 13 to 15 additionally, light is directed to the pad or patch structure 426 by a flexible fibreoptic 435 which has a

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436 arranged for mating connection connector corresponding connector 437 provided for the pad or patch structure. The pad or patch structure 426 has an imbedded fibreoptic arrangement 440 including a series of spaced lengths of fibreoptic 441 extending across the pad or patch structure. In the embodiment shown in figure 13, the fibreoptic path lengths form a serpentine arrangement, although a number of paths connected in parallel would also provide a convenient and workable arrangement. figure 14 the fibreoptic 440 includes .a shown in reflective surface cladding 442 surrounding a core 443. The surface cladding 442 ensures total internal reflection about the circumference of the fibreoptic except for an unclad window length 445 through which light can be emitted.

The pad or patch structure as shown in figures 12 to 15 provides a convenient arrangement in that light can be distributed widely over a large area of skin surface in a convenient and safe manner.

Particular benefits of the invention are the arrangement of the carrier and the photosensitive toxic material in discrete separate package portions, easily mixable at the required time in order to ensure that detrimental effects of early mixing or incorrect dosage ratios are avoided. The illumination apparatus configured to deliver non-laser light and/or configured to be placed against the target zone skin provide additional safety benefits and ensure that the skin is radiated to a consistent degree during treatment. This makes the kit particularly suitable for

consumer/home use.

Claims:

1. A cosmetic kit for use in inhibiting hair growth, the kit comprising:

> a) a marker substance including a photosensitive toxic material to tag subcutaneous hair growth cells of a target zone of skin; and,

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b) illumination apparatus for illuminating the skin in the target zone with light of a wavelength selected to effect preferential interaction with the hair growth cells tagged with the marker substance, resulting in a toxic effect inhibiting the hair growth capability of the tagged illuminated cells; wherein:

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(i) the marker substance is provided in a form comprising discrete portions of the photosensitive toxic material and a carrier material, the portions to be mixed prior to application to the target zone; and or,

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(ii) the illumination apparatus includes an applicator configured to deliver non-laser light and/or configured to be placed against the target zone skin to deliver the illumination to the tagged hair growth cells.

- 2. A cosmetic kit according to claim 1, wherein the portions of the photosensitive toxic material and the carrier material are dosed in a predetermined ratio.
- 3. A cosmetic kit according to claim 1 or claim 2, including a dispenser for dispensing the marker substance.
- A cosmetic kit according to any preceding claim,
 including a container for the carrier material and or the photosensitive toxic material.
- 5. A cosmetic kit according to claim 4, wherein the container affords shielding of the photosensitive toxic material.
 - 6. A cosmetic kit according to any preceding claim, wherein mixing means is provided for mixing the photosensitive toxic material and the carrier material.
 - 7. A cosmetic kit according to claim 6, wherein the mixing means permits mixing of the photosensitive toxic material and the carrier material at a zone substantially sealed from the user.
 - 8. A cosmetic kit according to any preceding claim, wherein a container for the photosensitive toxic material and the carrier material comprises respective sealed zones containing the photosensitive toxic material and the carrier material respectively,

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the zones being configured to be selectively breached or ruptured permitting mixing of the materials.

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- 9. A cosmetic kit according to claim 8, wherein a mixing zone is provided adjacent the breachable sealed zones, the mixing zone communicating with a dispensing outlet.
- 10. A cosmetic kit according to any preceding claim,
 wherein the illumination apparatus includes window
 means surface arranged to overlay the target zone
 skin, the illumination being directed via the window
 means.
- 11. A cosmetic kit according to any preceding claim, wherein the illumination apparatus includes an illumination applicator having a flexible structure arranged to conform to the target zone skin body part.

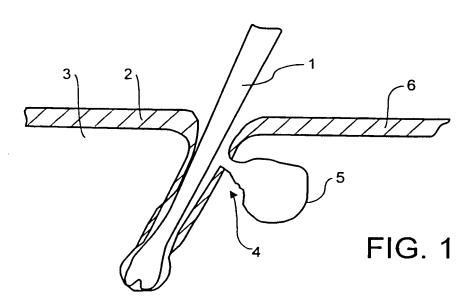
- 12. A cosmetic kit according to any preceding claim, wherein the apparatus includes:
- (a) an illumination applicator structure arranged to deliver illumination over a large area;
 - (b) a waveguide to connect with the illumination applicator; and,
- 30 (c) a light source to direct light along the waveguide.

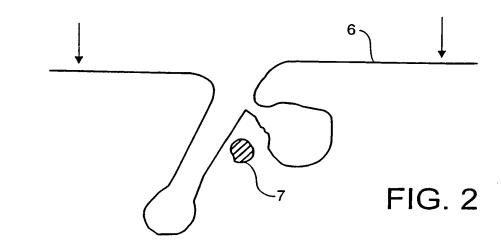
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- 13. A cosmetic kit according to any preceding claim, wherein the apparatus includes an illumination applicator structure having internal light reflecting walls and an associated light transmissive zone means through which reflected light is permitted to pass.
- 14. A cosmetic kit according to claim 13, wherein the applicator structure comprises one or more lengths of fibreoptic.
- 15. A cosmetic kit according to claim 13 or claim 14, wherein the applicator structure comprises a plurality of lengths of fibre optic, the lengths being arranged to deliver light at different zones of the applicator.



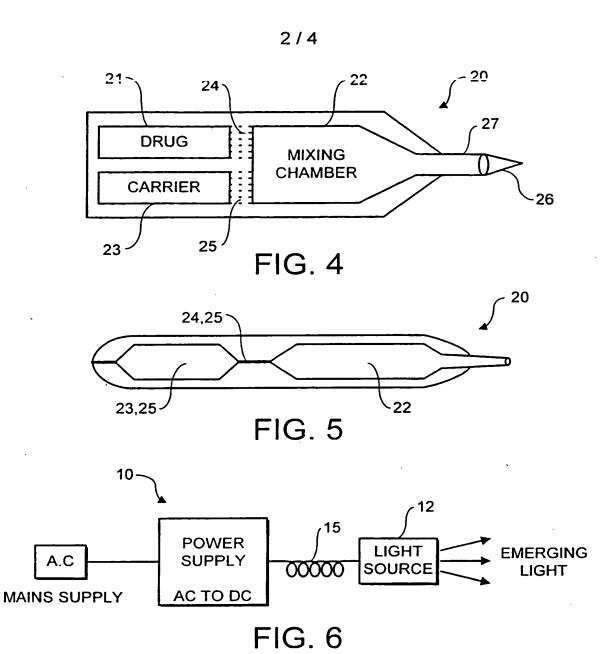


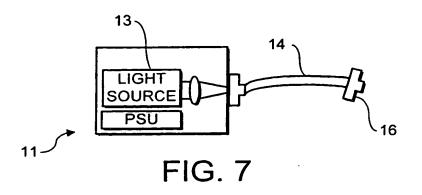




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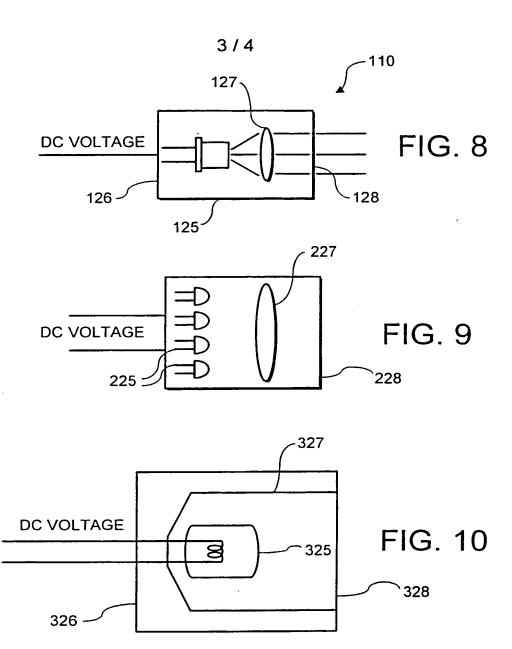
FIG. 3





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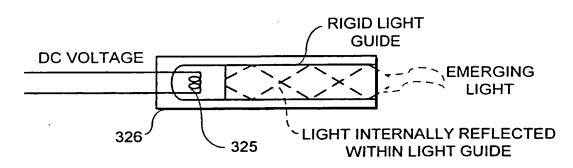
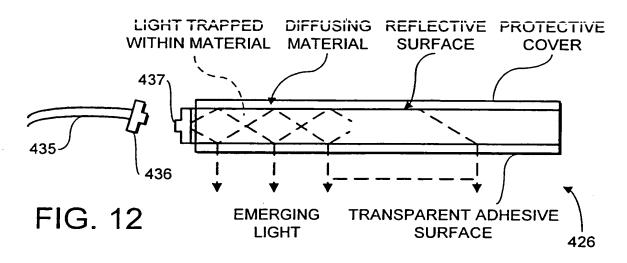
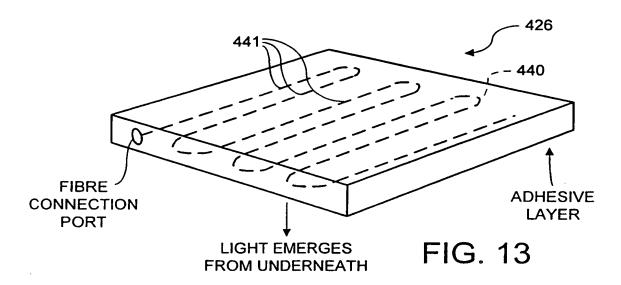


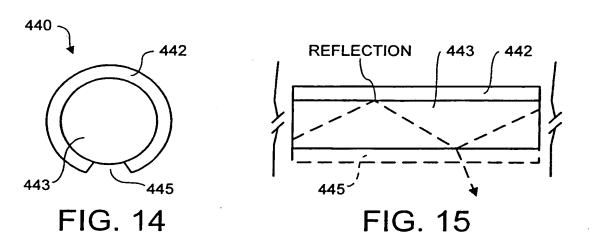
FIG. 11

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INTERNATIONAL SEARCH REPORT

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A. CLASSI IPC 7	FICATION OF SUBJECT MATTER A61N5/06 A61B18/18		
According to	o International Patent Classification (IPC) or to both national clas	sification and IPC	
B. FIELDS	SEARCHED		
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Documental	tion searched other than minimum documentation to the extent t	nat such documents are included in th	ne fields searched
Electronic d	ata base consulted during the international search (name of dat	a base and, where practical, search to	erms used)
EPO-In	ternal		
C. DOCUMI	ENTS CONSIDERED TO BE RELEVANT		
Category °	Citation of document, with indication, where appropriate, of the	e relevant passages	Relevant to claim No.
X	WO 97 32046 A (NEW YORK BLOOD (4 September 1997 (1997-09-04) page 3, line 13 - line 22 page 4, line 11 - line 28	CENTER INC)	1-7,10, 12
γ	page 5, line 3 - line 12		8,9,11,
•			13-15
X	WO 95 07077 A (NORWEGIAN RADIUM RESE ;DZIEGLEWSKA HANNA EVA (GI 16 March 1995 (1995-03-16) page 3, paragraph 1 page 5, paragraph 1 -page 6, pa page 8, paragraph 2 -page 10, page 10, p	1-7,10, 12	
		-/	
X Furth	ner documents are listed in the continuation of box C.	χ Patent family members	are listed in annex.
"A" docume	tegories of cited documents : ont defining the general state of the art which is not ered to be of particular relevance	"T" later document published afte or priority date and not in co cited to understand the princ invention	er the international filing date nflict with the application but siple or theory underlying the
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"O" docume other r	n or other special reason (as specified) ant referring to an oral disclosure, use, exhibition or neans	cannot be considered to invo document is combined with	nce; the claimed invention olive an inventive step when the one or more other such docu- ing obvious to a person skilled
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Date of the	actual completion of the international search	Date of mailing of the interna	ntional search report
19	9 September 2000	26/09/2000	
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INTERNALIONAL SEARCH REPORT

In. ational Application No
PCT/GB 00/02170

		PC1/GB 00/021/0
C.(Continue	ation) DOCUMENTS CONSIDERED TO BE RELEVANT	
Category °	Citation of document, with indication, where appropriate, of the relevant passages	Relevant to claim No.
X	US 5 669 916 A (ANDERSON RICHARD ROX) 23 September 1997 (1997-09-23) cited in the application column 4, line 25 - line 42	1,2
X	US 5 474 528 A (MESEROL PETER M) 12 December 1995 (1995-12-12) column 6, line 61 -column 7, line 9 column 8, line 17 -column 9, line 2	1-8, 10-12
Υ .	US 4 093 067 A (HOLLANDER JR EDWARD F) 6 June 1978 (1978-06-06) column 2, line 54 -column 3, line 2	8,9
Y	WO 99 10046 A (BIEL MERRILL A ;ADVANCED PHOTODYNAMIC TECHNOLO (US)) 4 March 1999 (1999-03-04) abstract	11
Y	US 5 849 027 A (CESATI CLAUDIO ET AL) 15 December 1998 (1998-12-15) column 4, line 20 - line 32 column 6, line 2 - line 13	13-15
A	US 3 736 933 A (SZABO B) 5 June 1973 (1973-06-05) abstract	8,9
A	US 5 835 648 A (ANDERSON STEVEN C ET AL) 10 November 1998 (1998-11-10) column 3, line 17 - line 59	13,14
A .	US 5 519 534 A (COLE JOHN ET AL) 21 May 1996 (1996-05-21) abstract	13
į		

1

Form PCT/ISA/210 (continuation of second sheet) (July 1992)

IN. RNATIONAL SEARCH REPORT

information on patent family members

Ir. ational Application No PCT/GB 00/02170

Patent document cited in search report		Publication date	Patent family member(s)	Publication date
WO 9732046	A	04-09-1997	AU 2135997 A CA 2247611 A EP 0883695 A JP 2000506140 T	16-09-1997 04-09-1997 16-12-1998 23-05-2000
WO 9507077	Α	16-03-1995	AU 7543794 A	27-03-1995
US 5669916	A .	23-09-1997	AU 689962 B AU 3685095 A CA 2201123 A EP 0783347 A JP 10506554 T WO 9609853 A US 5989267 A	09-04-1998 19-04-1996 04-04-1996 16-07-1997 30-06-1998 04-04-1996 23-11-1999
US 5474528	Α	12-12-1995	WO 9704836 A	13-02-1997
US 4093067	Α	06-06-1978	NONE	
WO 9910046	Α	04-03-1999	AU 9294298 A EP 1009483 A	16-03-1999 21-06-2000
US 5849027	Α	15-12-1998	NONE	
US 3736933	Α	05-06-1973	NONE	
US 5835648	Α	10-11-1998	NONE	
US 5519534	Α	21-05-1996	AU 2651795 A WO 9532441 A	18-12-1995 30-11-1995

Form PCT/ISA/210 (patent family annex) (July 1992)

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